

THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicant(s) : Beck et al.

Serial No. : RCE of 09/902,845

Cnfrm. No. : 9511

Filed : July 11, 2001

For : NOVEL 4-PHENYL SUBSTITUTED
TETRAHYDROISOQUINOLINES
THERAPEUTIC USE THEREOF

Examiner:
B. M. Robinson

Art Unit:
1625

**SECOND SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT
UNDER 37 CFR §§ 1.97-1.98**

Mail Stop RCE

Commissioner for Patents
P.O. Box 1450
Alexandria, VA 22313-1450

Dear Sir:

Pursuant to 37 CFR §§ 1.97-1.98, applicants hereby bring to the attention of the United States Patent and Trademark Office, the enclosed references listed on the attached PTO-1449 form. Document Nos. 158, 160, 183, and 212 were cited in the August 29, 2003, Communication Pursuant to Article 96(2) EPC (“European Communication”) issued in the corresponding European application. A copy of the European Communication is enclosed.

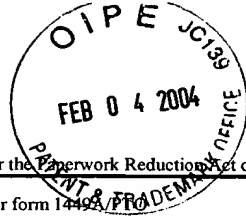
Respectfully submitted,

Date: February 2, 2004

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INFORMATION DISCLOSURE STATEMENT BY APPLICANT (use as many sheets as necessary)				Complete if Known	
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				Filing Date	July 11, 2001
				First Named Inventor	Beck et al.
				Art Unit	1625
				Examiner Name	B. M. Robinson
Sheet	1	of	9	Attorney Docket Number	20011/1381

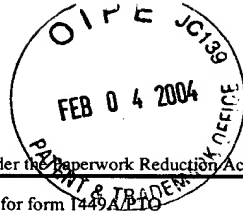
U.S. PATENT DOCUMENTS					
Examiner Initials*	Cite No. ¹	U.S. Patent Document	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
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		Country Code ³ Number ⁴ Kind Code ⁵ (if known)				
	1 /	EP 0 599 338 A2	06/01/94	Europe		
	2 /	CH 538 477	08/05/73	Switzerland		
	3 /	JP 04193867	07/13/92	Japan		
	4 /	DE 2 062 001	07/15/71	Germany		
	5 /	WO 97/36876	10/09/97	WIPO		
	6 /	WO 97/23458	07/03/97	WIPO		
	7 /	WO 90/05525	05/31/90	WIPO		
	8 /	WO 90/05729	05/31/90	WIPO		
	9 /	WO 94/02461	02/03/94	WIPO		
	10 /	WO 94/02595	02/03/94	WIPO		
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	14 /	WO 94/04496	03/03/94	WIPO		
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	19 /	WO 94/10167	05/11/94	WIPO		
	20 /	WO 94/10168	05/11/94	WIPO		
	21 /	WO 94/10170	05/11/94	WIPO		
	22 /	WO 94/11368	05/26/94	WIPO		
	23 /	WO 94/13639	06/23/94	WIPO		
	24 /	WO 94/13663	06/23/94	WIPO		
	25 /	WO 94/14767	07/07/94	WIPO		
	26 /	WO 94/15903	07/21/94	WIPO		
	27 /	WO 94/19320	09/01/94	WIPO		
	28 /	WO 94/19323	09/01/94	WIPO		
	29 /	WO 94/20500	09/15/94	WIPO		
	30 /	WO 91/09844	07/11/91	WIPO		

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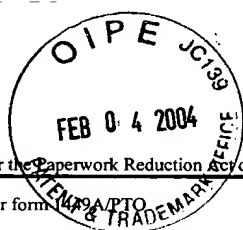
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	31 ✓	WO 91/18899	12/12/91	WIPO		
	32 ✓	WO 92/01688	02/06/92	WIPO		
	33 ✓	WO 92/06079	04/16/92	WIPO		
	34 ✓	WO 92/12151	07/23/92	WIPO		
	35 ✓	WO 92/15585	09/17/92	WIPO		
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	38 ✓	WO 92/20676	11/26/92	WIPO		
	39 ✓	WO 92/21677	12/10/92	WIPO		
	40 ✓	WO 92/22569	12/23/92	WIPO		
	41 ✓	WO 93/00330	01/07/93	WIPO		
	42 ✓	WO 93/00331	01/07/93	WIPO		
	43 ✓	WO 93/01159	01/21/93	WIPO		
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	49 ✓	WO 93/10073	05/27/93	WIPO		
	50 ✓	WO 93/14084	07/22/93	WIPO		
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	52 ✓	WO 93/18023	09/16/93	WIPO		
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	57 ✓	WO 93/24465	12/09/93	WIPO		
	58 ✓	WO 94/00440	01/06/94	WIPO		
	59 ✓	WO 94/01402	01/20/94	WIPO		
	60 ✓	WO 94/26735	11/24/94	WIPO		

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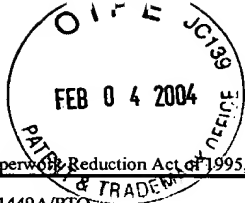
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	61 ✓	WO 94/26740	11/24/94	WIPO		
	62 ✓	WO 94/29309	12/22/94	WIPO		
	63 ✓	WO 95/02595	01/26/95	WIPO		
	64 ✓	WO 95/04040	02/09/95	WIPO		
	65 ✓	WO 95/04042	02/09/95	WIPO		
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	69 ✓	WO 95/08549	03/30/95	WIPO		
	70 ✓	WO 95/11880	05/04/95	WIPO		
	71 ✓	WO 95/14017	05/26/95	WIPO		
	72 ✓	WO 95/15311	06/08/95	WIPO		
	73 ✓	WO 95/16679	06/22/95	WIPO		
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	79 ✓	WO 95/22525	08/24/95	WIPO		
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	81 ✓	WO 95/26338	10/05/95	WIPO		
	82 ✓	WO 95/28418	10/26/95	WIPO		
	83 ✓	WO 95/30674	11/16/95	WIPO		
	84 ✓	WO 95/30687	11/16/95	WIPO		
	85 ✓	WO 95/33744	12/14/95	WIPO		
	86 ✓	WO 96/05181	02/22/96	WIPO		
	87 ✓	WO 96/05193	02/22/96	WIPO		
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	89 ✓	WO 96/06094	02/29/96	WIPO		
	90 ✓	WO 96/07649	03/14/96	WIPO		

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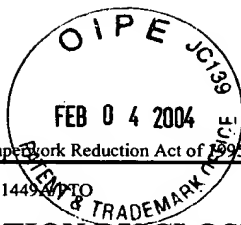
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	91	WO 96/10562	04/11/96	WIPO		
	92	WO 96/16939	06/06/96	WIPO		
	93	WO 96/18643	06/20/96	WIPO		
	94	WO 96/20197	07/04/96	WIPO		
	95	WO 96/21661	07/18/96	WIPO		
	96	WO 96/29304	09/26/96	WIPO		
	97	WO 96/29317	09/26/96	WIPO		
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	99	WO 96/29328	09/26/96	WIPO		
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	106	WO 97/08144	03/06/97	WIPO		
	107	WO 97/14671	04/24/97	WIPO		
	108	WO 97/17362	05/15/97	WIPO		
	109	WO 97/18206	05/22/97	WIPO		
	110	WO 97/19084	05/29/97	WIPO		
	111	WO 97/19942	06/05/97	WIPO		
	112	WO 97/21702	06/19/97	WIPO		
	113	WO 97/49710	12/31/97	WIPO		
	114	GB 2 266 529 A	11/03/93	Great Britain		
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	116	GB 2 269 170 A	02/02/94	Great Britain		
	117	GB 2 269 590 A	02/16/94	Great Britain		
	118	GB 2 271 774 A	04/27/94	Great Britain		
	119	GB 2 292 144 A	02/14/96	Great Britain		
	120	GB 2 293 168 A	03/20/96	Great Britain		

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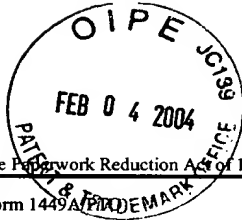
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	121	GB 2 293 169 A	03/20/96	Great Britain		
	122	GB 2 302 689 A	01/29/97	Great Britain		
	123	EP 0 360 390 A1	03/28/90	Europe		
	124	EP 0 517 589 B1	12/27/96	Europe		
	125	EP 0 520 555 A1	12/30/92	Europe		
	126	EP 0 520 555 B1	09/01/99	Europe		
	127	EP 0 522 808 A2	01/13/93	Europe		
	128	EP 0 528 495 A1	02/24/93	Europe		
	129	EP 0 532 456 B1	03/29/95	Europe		
	130	EP 0 533 280 B1	03/24/93	Europe		
	131	EP 0 536 817 A1	04/14/93	Europe		
	132	EP 0 545 478 A1	06/09/93	Europe		
	133	EP 0 558 156 A2	09/01/93	Europe		
	134	EP 0 577 394 B1	01/05/94	Europe		
	135	EP 0 585 913 B1	03/09/94	Europe		
	136	EP 0 599 538 A1	06/01/94	Europe		
	137	EP 0 610 793 A1	08/17/94	Europe		
	138	EP 0 634 402 A1	01/18/95	Europe		
	139	EP 0 686 629 A2	12/13/95	Europe		
	140	EP 0 693 489 A1	01/24/96	Europe		
	141	EP 0 694 535 A1	01/31/96	Europe		
	142	EP 0 699 655 B1	09/24/97	Europe		
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	145	EP 0 429 366 B1	05/29/91	Europe		
	146	EP 0 430 771 B1	06/05/91	Europe		
	147	EP 0 436 334 B1	07/10/91	Europe		
	148	EP 0 443 132 B1	08/28/91	Europe		
	149	EP 0 482 539 B1	04/29/92	Europe		
	150	EP 0 498 069 B1	08/12/92	Europe		

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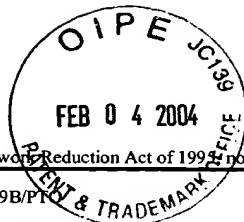
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	151	US-5,532,244	07/02/96	Wong et al.	
	152	US-5,444,070	08/22/95	Moldt et al.	
	153	US-6,136,803	10/24/00	Freedman et al.	
	154	US-5,789,449	08/04/98	Norden	
	155	US-4,902,710	02/20/90	Foster et al.	
	156	US-6,121,261	09/19/00	Glatt et al.	
	157	US-4,843,071	06/27/89	Hohenwarter	
	158	US-3,666,763	05/30/72	Grethe et al.	
	159	US-5,654,296	08/05/97	Kato et al.	
	160	US-4,564,613	01/14/86	Boltze et al.	

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	161	EP 0 499 313 B1	08/19/92	Europe		
	162	EP 0 512 901 B1	11/11/92	Europe		
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	168	EP 0 515 681 A1	12/02/92	Europe		
	169	EP 0 699 674 A1	03/06/96	Europe		
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	181	EP 0 400 319 A1	12/05/90	Europe		
	182	WO 02/04455 A3	01/17/02	WIPO		
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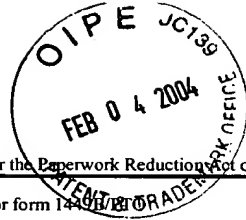
Substitute for form 1449B/PTO INFORMATION DISCLOSURE STATEMENT BY APPLICANT (use as many sheets as necessary)		Complete if Known		
		Application Number	RCE of 09/902,845	
		Filing Date	July 11, 2001	
		First Named Inventor	Beck et al.	
		Group Art Unit	1625	
		Examiner Name	B. M. Robinson	
Sheet	7	9	Attorney Docket Number	20011/1381

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	184 ✓	Cliffe et al., "(S)-N-tert-Butyl-3-(4-(2-methoxyphenyl)-piperazin-1-yl)-2-phenylpropanamide [(S)-WAY- 100135]: A Selective Antagonist at Presynaptic and Postsynaptic-5-HT _{1A} Receptors," <u>J. Med. Chem.</u> 36:1509-10 (1993)	
	185 ✓	Salama et al., "Antigenic Determinants Responsible for the Reactions of Drug-Dependent Antibodies with Blood Cells," <u>British Journal of Haematology</u> 78:535-539 (1991)	
	186 ✓	Trepanier et al., "3,4-Dihydroisocarbostyrl and 1,2,3,4-Tetrahydroisoquinoline Derivatives of Ephedrine," <u>Journal of Medicinal Chemistry</u> 16(4):342-347 (1973)	
	187 ✓	Miller et al., "An Efficient Synthesis of 4-Aryl-1,2,3,4-Tetrahydroisoquinolines," <u>Synthetic Communications</u> 24(8):1187-1193 (1994)	
	188 ✓	Tirelli et al., "Differential Effects of Direct and Indirect Dopamine Agonists on the Induction of Gnawing in C57BI/6J Mice," <u>Journal of Pharmacology and Experimental Therapeutics</u> 273(1):7-15 (1995)	
	189 ✓	Jacob et al., "Dopamine Agonist Properties of N-Alkyl-4-(3,4-dihydroxyphenyl)-1,2,3,4-tetrahydroisoquinolines," <u>J. Med. Chem.</u> 24:1013-1015 (1981)	
	190 ✓	Ishikura et al., "The Synthesis of 4-Substituted Isoquinoline Derivatives from Diethyl (4-Isoquinolyl) Borane," <u>Heterocycles</u> 26:1603-1610 (1987)	
	191 ✓	Berge et al., "Pharmaceutical Salts," <u>Journal of Pharmaceutical Sciences</u> 66(1):1-19 (1977)	
	192 ✓	Bundgaard, <u>Design of Prodrugs</u> , Amsterdam, The Netherlands: Elsevier Science Publishers B.V. (1985) (Table of Contents only)	
	193 ✓	Krogsgaard-Larsen et al., eds., <u>A Textbook of Drug Design and Development</u> , Chur, Switzerland: Harwood Academic Publishers GmbH (1991) (portion of Table of Contents only)	

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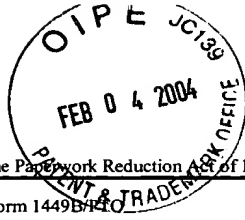
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	194 ✓	Bundgaard, "Means to Enhance Penetration," <u>Advanced Drug Delivery Reviews</u> 8:1-38 (1992)	
	195 ✓	Nielsen et al., "Glycolamide Esters as Biolabile Prodrugs of Carboxylic Acid Agents: Synthesis, Stability, Bioconversion, and Physicochemical Properties," <u>Journal of Pharmaceutical Sciences</u> 77(4):285-298 (1988)	
	196 ✓	Kakeya et al., "Studies on Prodrugs of Cephalosporins. I. Synthesis and Biological Properties of Glycyloxybenzoyloxymethyl and Glycylaminobenzoyloxymethyl Esters of 7β-[2-(2-Aminothiazol-4-yl)-(Z)-2-Methoxyiminoacetamido]-3-Methyl-3-Cephem-4-Carboxylic Acid," <u>Chem. Pharm. Bull.</u> 32(2):692-698 (1984)	
	197 ✓	Middlemiss et al., "Centrally Active 5-HT Receptor Agonists and Antagonists," <u>Neuroscience & Biobehavioral Reviews</u> 16:75-82 (1992)	
	198 ✓	Greene et al., <u>Protective Groups in Organic Synthesis</u> , 2d. Ed., New York, New York: John Wiley & Sons, Inc. (1991) (Table of Contents only)	
	199 ✓	McOmie, ed., <u>Protective Groups in Organic Chemistry</u> , London: Plenum Press (1973) (Table of Contents only)	
	200 ✓	Jorgenson, "Preparation of Ketones from the Reaction of Organolithium Reagents with Carboxylic Acids," Dauben et al., eds., <u>Organic Reactions</u> , Vol. 18, New York, New York: John Wiley & Sons, Inc., Chapter 1 (1970) (Table of Contents only)	
	201 ✓	Larock, <u>Comprehensive Organic Transformations: A Guide to Functional Group Preparation</u> , New York, New York: VCH Publishers, Inc. (1989) (Table of Contents only)	
	202 ✓	Venkov et al., "A New Synthesis of 1,2,3,4-Tetrahydro-2-Methyl-4-Phenylisoquinolines," <u>Synthesis</u> 253-255 (1990)	
	203 ✓	Dandridge et al., "Synthesis, Resolution, Absolute Stereochemistry, and Enantioselectivity of 3', 4'- Dihydroxynomifensine," <u>J. Med. Chem.</u> 27:28-35 (1984)	
	204 ✓	Kihara et al., "A Convenient Synthesis of 4-Substituted 1,2,3,4-Tetrahydroisoquinolin-4-OLS by a Novel Intramolecular Barbier Reaction and by an Insertion Reaction: Reaction Scope and Limitations," <u>Tetrahedron</u> 48(1): 67-78 (1992)	

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	205 ✓	Hudlicky, "Fluorination with Diethylaminosulfur Trifluoride and Related Aminofluorosulfuranes," <u>Organic Reactions</u> 35:513-637 (1985)	
	206 ✓	Gao et al., "Asymmetric Hetero Diels-Alder Reaction Catalyzed by Stable and Easily Prepared CAB Catalysts," <u>Tetrahedron</u> 50(4):979-988 (1994)	
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	208 ✓	Stille, "Zur pharmakologischen Prufung von Antidepressiva am Beispiel eines Dibenzodiazepins," <u>Arzneimittel-Forschung</u> 14:534-537 (1964) (English summary included)	
	209 ✓	Blomberg et al., "The Barbier Reaction – A One Step Alternative for Syntheses via Organomagnesium Compounds," <u>Synthesis</u> pp. 18-30 (1977)	
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	211 ✓	Kihara et al., "Synthesis and Pharmacological Evaluation of Phenolic 2-Methyl-4-Phenyl-1,2,3,4,- Tetrahydroisoquinolin-4-ols As New Norepinephrine Potentiator," <u>Drug Design and Discovery</u> 11(3):175-183 (1994)	
	212 ✓	Kihara et al., "Synthesis and Enantioselectivity of Optically Active 1- and 3-Substituted 4-Phenyl-1,2,3,4- Tetrahydroisoquinolin-4-ols and Related Compounds As Norepinephrine Potentiators," <u>Chemical and Pharmaceutical Bulletin</u> 43(9):1543-1546 (1995)	
	213 ✓	Aihara et al., "Increasing 5-Lipoxygenase Inhibitory Activities by Oxidative Conversion of <i>o</i> -Methoxyphenols to Catechols Using a Cu ²⁺ -Ascorbic Acid-O ₂ System," <u>Chem. Pharm. Bull.</u> 38(3):842-844 (1990)	

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